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IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Appl. No. : 10/758,415  
Applicant : William S. Brusilow  
Filed : January 6, 2004  
TC/A.U. : 1614  
Examiner :

Docket No. : 2930-109  
Customer No. : 06449  
Confirmation No. : 5654

**INFORMATION DISCLOSURE STATEMENT**

Director of the United States Patent  
and Trademark Office  
P.O. Box 1450  
Alexandria, Virginia 22313-1450

Dear Sir:


Under the provisions of 37 C.F.R. §§ 1.56, 1.97 and 1.98, Applicant submits herewith information that the Office may wish to consider in examination of the subject application. Materials submitted for consideration are listed on the attached form PTO-1449.

Respectfully submitted,

By

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RBM/cb

<b>INFORMATION DISCLOSURE STATEMENT OF APPLICANT</b> 				<i>Complete if Known</i>	
				Application Number	10/758,415
				Filing Date	January 16, 2004
				First Named Inventor	William S. Brusilow
				Group Art Unit	1614
Examiner Name					
Sheet	1	of	2	Attorney Docket Number	2930-109
<b>OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS</b>					
Examiner Initials*	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published			T <sup>2</sup>
	AA	Blei, Andres T., et al., "Ammonia-Induced Brain Edema and Intracranial Hypertension in Rats After Portacaval Anastomosis," <i>Hepatology</i> 19(6): 1437-1444, June 1994.			
	AB	Brusilow, Saul W. "Inborn Errors of Urea Synthesis," In: Scriver CR, Lloyd JK, eds. <u>Genetic and Metabolic Disease in Pediatrics</u> . 5: 140-165, London: Butterworths, 1985.			
	AC	Brusilow, Saul W., et al., "Urea Cycle Disorders: Diagnosis, Pathophysiology, and Therapy," <i>Advances in Pediatrics</i> 43:127-170, 1996.			
	AD	Butterworth, R.F., "Effects of Hyperammonaemia on Brain Function," <i>J. Inher. Metab. Dis.</i> 21(1):6-20, 1998.			
	AE	Cordoba, Juan, et al., "Brain Edema and Hepatic Encephalopathy," <i>Seminars in Liver Disease</i> 16(3): 271-280, 1996.			
	AF	Folbergrova, J., "Free Glutamine Level in the Rat Brain In Vivo After Methionine Sulphoximine Administration," <i>Physiologia Bohemoslovenica</i> 13:21-26, 1963.			
	AG	Gershoff, S.N., et al., "The Relative Effect of Methionine Sulfoximine on Different Animal Species," <i>J. Nutr.</i> 45:451-458, 1951.			
	AH	Häussinger, Dieter, et al., "Pathogenesis of Hepatic Encephalopathy," <i>Journal of Gastroenterology and Hepatology</i> 17:S256-S259, 2002.			
	AI	Hawkins, Richard, et al., "Effect of Reducing Brain Glutamine Synthesis on Metabolic Symptoms of Hepatic Encephalopathy," <i>Journal of Neurochemistry</i> 60(3):1000-1006, 1993.			
	AJ	Hawkins, Richard, et al., "Hyperammonaemia Does Not Impair Brain Function in the Absence of Net Glutamine Synthesis," <i>Biochem. J.</i> 277:697-703, 1991.			
	AK	Hirata, Takahiko, et al., "Impaired Pial Arteriolar Reactivity to Hypercapnia During Hyperammonemia Depends on Glutamine Synthesis," <i>Stroke</i> 27(4): 729-736, 1996.			
	AL	Jonung, Torbjorn, et al., "Methionine Sulfoximine Prevents the Accumulation of Large Neutral Amino Acids in Brain of Hyperammonemic Rats," <i>J. Surgical Research</i> 36:349-353, 1984.			
	AM	Krakoff, Irwin H., et al., "Effect of Methionine Sulfoximine in Man," <i>J. Pharm. Experimental Ther.</i> 2:599-604, 1961.			
	AN	Lamar, C., et al., "The Duration of the Inhibition of Glutamine Synthetase by Methionine Sulfoximine," <i>Biochemical Pharmacology</i> 17:636-642, 1968.			
Examiner Signature				Date Considered	

\*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. <sup>1</sup>Unique citation designation number. <sup>2</sup>Applicant is to place a check mark here if English language Translation is attached.

**INFORMATION DISCLOSURE  
STATEMENT BY APPLICANT**



Complete if Known

Application Number	10/758,415
Filing Date	January 16, 2004
First Named Inventor	William S. Brusilow
Group Art Unit	1614
Examiner Name	

Sheet	2	OF	2	Attorney Docket Number	2930-109
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**OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS**

Examiner Initials*	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published	T <sup>2</sup>
	AO	Master, Sonali, et al., "Cerebral Blood Flow and the Development of Ammonia-Induced Brain Edema in Rats After Portacaval Anastomosis," <i>Hepatology</i> 30(4): 876-880, 1999.	
	AP	Norenberg, Michael D., et al., "Fine Structural Localization of Glutamine Synthetase in Astrocytes of Rat Brain," <i>Brain Research</i> 161:303-310, 1979.	
	AQ	Richman, Paul G., et al., "Inhibition of $\gamma$ -Glutamylcystein Synthetase by L-Methionine-S-Sulfoximine," <i>J. Biological Chemistry</i> 248(19): 6684-6690, 1973.	
	AR	Rowe, W. Bruce, et al., "Identification of L-Methionine-S-Sulfoximine as the Convulsant Isomer of Methionine Sulfoximine," <i>Proceedings of the National Academy of Sciences</i> 66(2): 500-506, June 1970.	
	AS	Sellinger, Otto Z., et al., "Methionine Sulfoximine Seizures. VII. the Dissociation of the Convulsant and Glutamine Synthetase Inhibitory Effects," <i>J. Pharmacology &amp; Experimental Therapeutics</i> 161(1): 212-222, 1968.	
	AT	Sugimoto, Hideyoshi, et al., "Methionine Sulfoximine, A Glutamine Synthetase Inhibitor, Attenuates Increased Extracellular Potassium Activity During Acute Hyperammonemia," <i>Journal of Cerebral Blood Flow &amp; Metabolism</i> , 17:44-49, 1997.	
	AU	Takahashi, Hideo, et al., "Inhibition of Brain Glutamine Accumulation Prevents Cerebral Edema in Hyperammonemic Rats," <i>American Physiological Society</i> 261:H825-H829, 1991.	
	AV	Voorhies, Theresa M., "Acute Hyperammonemia in the Young Primate: Physiologic and Neuropathologic Correlates," <i>Pediatric Research</i> 17(12):970-975, 1983.	
	AW	Wada, Juhn A., et al., "The Susceptibility of Auditory Stimuli of Animals Treated with Methionine Sulfoximine," <i>Experimental Neurology</i> 15:157-165, 196.	
	AX	Warren, Kenneth S., et al., "Effect of an Inhibitor of Glutamine Synthesis (Methionine Sulfoximine) on Ammonia Toxicity and Metabolism," <i>J. Lab. &amp; Clin. Med.</i> 64(3): 442-449, 1964.	
	AY	Watson, Alan J., et al. "Transient Idiopathic Hyperammonaemia in Adults," <i>The Lancet</i> 1271-1274, December 7, 1985.	
	AZ	Willard-Mack, C.L., et al., "Inhibition of Glutamine Synthetase Reduces Ammonia-Induced Astrocyte Swelling in Rat," <i>Neuroscience</i> 71(2): 589-599, 1996.	
	BA	Zwingmann, Claudia, et al., "Multinuclear NMR Spectroscopy Studies on NH <sub>4</sub> Cl-Induced Metabolic Alterations and Detoxification Processes in Primary Astrocytes and Glioma Cells," <i>Dev. Neurosci</i> 20:417-426, 1998.	
	BB	Apostolakis et al., <i>Brain Research Bulletin</i> , Vol. 23, pp. 257-262 (1989).	
	BC	Ginefri-Gayet et al., <i>Pharmacology Biochemistry and Behavior</i> , Vol. 43, pp. 173-179 (1992).	
	BD	Takahashi et al., <i>Circulation Research</i> , 71(5), 1220-1230 (November, 1992).	
	BE	Harth et al., <i>J. Exp. Med.</i> , 189(9), 1425-1435 (May 3, 1999).	
Examiner Signature		Date Considered	

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